

TECH CENTER 1600/2900

RAW SEQUENCE LISTING

4 <110> APPLICANT: Nicolaides, Nicholas

PATENT APPLICATION: US/09/749,601A

DATE: 03/18/2002 TIME: 14:45:17

Input Set : A:\00069.JHU.SEQ.TXT

Output Set: N:\CRF3\03182002\1749601A.raw

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5
         Grasso, Luigi
 6
         Sass, Philip
         Kinzler, Kenneth
         Vogelstein, Bert
10 <120> TITLE OF INVENTION: A method for generating hypermutable
11
         plants
13 <130> FILE REFERENCE: 01107.00069
15 <140> CURRENT APPLICATION NUMBER: 09/749,601A
16 <141> CURRENT FILING DATE: 2000-12-28
18 <150> PRIOR APPLICATION NUMBER: 60/183,333
19 <151> PRIOR FILING DATE: 2000-02-18
21 <160> NUMBER OF SEQ ID NOS: 14
23 <170> SOFTWARE: FastSEQ for Windows Version 4.0
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26 <211> LENGTH: 17
27 <212> TYPE: DNA
28 <213> ORGANISM: Human sapiens
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                                                                           120
49 gagttagtag aaaacagtet ggatgetggt gecaetaata ttgatetaaa gettaaggae
                                                                           180
50 tatggagtgg atcttattga agtttcagac aatggatgtg gggtagaaga agaaaacttc
                                                                           240
51 gaaggettaa etetgaaaca teacacatet aagatteaag agtttgeega eetaaeteag
                                                                           300
52 gttgaaactt ttggctttcg gggggaagct ctgagctcac tttgtgcact gagcgatgtc
                                                                           360
53 accatttcta cctgccacgc atcggcgaag gttggaactc gactgatgtt tgatcacaat
                                                                           420
54 gggaaaatta teeagaaaac eeectaeeee egeeeeagag ggaeeacagt cagegtgeag
                                                                           480
55 cagttatttt ccacactacc tgtgcgccat aaggaatttc aaaggaatat taagaaggag
                                                                           540
56 tatgccaaaa tggtccaggt cttacatgca tactgtatca tttcagcagg catccgtgta
                                                                           600
                                                                           660
57 agttgcacca atcagcttgg acaaggaaaa cgacagcctg tggtatgcac aggtggaagc
58 cccagcataa aggaaaatat cggctctgtg tttgggcaga agcagttgca aagcctcatt
                                                                           720
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RAW SEQUENCE LISTING

PATENT APPLICATION: US/09/749,601A

PECE/VED

DATE: 03/18/2002 MAY I 5 2002

TIME: 14:4 FEH CENTER 1600/2900

Input Set : A:\00069.JHU.SEQ.TXT

Output Set: N:\CRF3\03182002\I749601A.raw

| 59 | ccttttgttc | agctgccccc | tagtgactcc | gtgtgtgaag | agtacggttt | gagctgttcg | 780 |
|-----|-------------|--------------|-------------|------------|------------|--------------|-------|
| 60 | gatgctctgc | ataatctttt | ttacatctca | ggtttcattt | cacaatgcac | gcatggagtt | 840 |
| 61 | ggaaggagtt | caacagacag | acagtttttc | tttatcaacc | ggcggccttg | tgacccagca | 900 |
| 62 | aaggtctgca | gactcgtgaa | tgaggtctac | cacatgtata | atcgacacca | gtatccattt | 960 |
| | | | | | tcaatgttac | | 1020 |
| 64 | aggcaaattt | tgctacaaga | ggaaaagctt | ttgttggcag | ttttaaagac | ctctttgata | 1080 |
| 65 | ggaatgtttg | atagtgatgt | caacaagcta | aatgtcagtc | agcagccact | gctggatgtt | 1140 |
| 66 | gaaggtaact | taataaaaat | gcatgcagcg | gatttggaaa | agcccatggt | agaaaagcag | 1200 |
| 67 | gatcaatccc | cttcattaag | gactggagaa | gaaaaaaaag | acgtgtccat | ttccagactg | 1260 |
| 68 | cgagaggcct | tttctcttcg | tcacacaaca | gagaacaagc | ctcacagccc | aaagactcca | 1320 |
| 69 | gaaccaagaa | ggagccctct | aggacagaaa | aggggtatgc | tgtcttctag | cacttcaggt | 1380 |
| 70 | gccatctctg | acaaaggcgt | cctgagacct | cagaaagagg | cagtgagttc | cagtcacgga | 1.440 |
| 71 | cccagtgacc | ctacggacag | agcggaggtg | gagaaggact | cggggcacgg | cagcacttcc | 1500 |
| 72 | gtggattctg | aggggttcag | catcccagac | acgggcagtc | actgcagcag | cgagtatgcg | 1560 |
| 73 | gccagctccc | caggggacag | gggctcgcag | gaacatgtgg | actctcagga | gaaagcgcct | 1620 |
| 74 | gaaactgacg | actctttttc | agatgtggac | tgccattcaa | accaggaaga | taccggatgt | 1680 |
| 75 | aaatttcgag | ttttgcctca | gccaactaat | ctcgcaaccc | caaacacaaa | gcgttttaaa | 1740 |
| 76 | aaagaagaaa | ttctttccag | ttctgacatt | tgtcaaaagt | tagtaaatac | tcaggacatg | 1800 |
| 77 | tcagcctctc | aggttgatgt | agctgtgaaa | attaataaga | aagttgtgcc | cctggacttt | 1860 |
| 78 | tctatgagtt | ctttagctaa | acgaataaag | cagttacatc | atgaagcaca | gcaaagtgaa | 1920 |
| 79 | ggggaacaga | attacaggaa | gtttagggca | aagatttgtc | ctggagaaaa | tcaagcagcc | 1980 |
| 80 | gaagatgaac | taagaaaaga | gataagtaaa | acgatgtttg | cagaaatgga | aatcattggt | 2040 |
| 81 | cagtttaacc | tgggatttat | aataaccaaa | ctgaatgagg | atatcttcat | agtggaccag | 2100 |
| 82 | catgccacgg | acgagaagta | taacttcgag | atgctgcagc | agcacaccgt | gctccagggg | 2160 |
| | | | | | ttaatgaagc | | 2220 |
| 84 | gaaaatctgg | aaatatttag | aaagaatggc | tttgattttg | ttatcgatga | aaatgctcca | 2280 |
| | | | | | aaaactggac | | 2340 |
| | | | | | ctggggtcat | | 2400 |
| | | | | | agtcggtgat | | 2460 |
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| | | | | | ccaacctggg | | 2580 |
| | cagaactag | | | _ | | | 2589 |
| 92 | <210> SEQ 1 | ID NO: 4 | | | | | |
| 93 | <211> LENGT | TH: 2340 | | | | | |
| 94 | <212> TYPE: | DNA | | | | | |
| 95 | <213> ORGAN | NISM: Arabio | opsis thali | ana | | | |
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| 98 | atgcaaggag | attcttctcc | gtctccgacg | actactaget | ctcctttgat | aagacctata | 60 |
| | | | | | tcttagacct | | 120 |
| | | | | | | gattaacctc | 180 |
| | | | | | | ttccccaacc | 240 |
| | | | | | | acttaagcat | 300 |
| | | | | | | ı tggttttaga | 360 |
| | | | | | | aagaacaaag | 420 |
| | | | | | | tgctgaaaag | 480 |
| | | | | | | taatttacct | 540 |
| | | | _ | | | tgtatctta | 600 |
| | | | | | | cacgactggg | 660 |
| 109 | aaaacccaa | ı agtctgttgt | gctgaacaca | caagggaggg | gttcacttaa | agataatatc | 720. |
| | | | | | | | |

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Output Set: N:\CRF3\03182002\I749601A.raw

| 110 | ataacagttt | tcggcattag | tacctttaca | agtctacagc | ctggtactgg | acgcaattta | 780 | | | | | | |
|-----|--|---|------------|--|--------------------------|------------|--------------|--|--|--|--|--|--|
| | | _ | | | atatgccaaa | | 840 | | | | | | |
| 112 | ttggtgaatg | agttatataa | agatacaagt | tctcggaaat | atccagttac | cattctggat | 900 | | | | | | |
| | | | | | ccgataaaag | | 960 | | | | | | |
| 114 | ttttctgacg | agacttctgt | tatcggttct | ttgagggaag | gtctgaacga | gatatattcc | 1020 | | | | | | |
| | | | | | attcggagca | | 1080 | | | | | | |
| | | | | - | cagaagggat | | 1140 | | | | | | |
| | | | | | | | 1200 1260 | | | | | | |
| | l7 gtcagttcta aaacaagact aggggaagct attgagaaag aaaatccatc cttaaggg l8 gttgaaattg ataatagttc gccaatggag aagtttaagt ttgagatcaa ggcatgtg l9 acgaagaaag gggaaggttc tttatcagtc catgatgtaa ctcaccttga caagacac | | | | | | | | | | | | |
| | | | 1320 | | | | | | | | | | |
| | | | | ngcaag taaagacttg 13 naccat gggaaaaaga 14 | | | | | | | | | |
| | | geageeget etagetttge ceagteaact ttgaataett ttgttaceat gggaaaaag | | | | | | | | | | | |
| | _ | _ | | | 1500 | | | | | | | | |
| | | | _ | | tagcttcaag | | 1560 | | | | | | |
| | | | | | atatgacacc | | 1620 | | | | | | |
| 125 | gattctgaac | taggcaatcg | gatttctcct | ggaacacaag | ctgataatgt | tgaaagacat | 1680 | | | | | | |
| 126 | gagagagtac | tcgggcaatt | caatcttggg | ttcatcattg | caaaattgga | gcgagatctg | 1740 | | | | | | |
| 127 | ttcattgtgg | atcagcatgc | agctgatgag | aaattcaact | tcgaacattt | agcaaggtca | 1800 | | | | | | |
| | | - | _ | _ | tggaactctc | | 1860 | | | | | | |
| | | | | | atggctttct | | 1920 | | | | | | |
| | | | | | ccattcctta | | 1980 | | | | | | |
| | | | | | ctctaggaga | | 2040 | | | | | | |
| | | | | | attcgatttg | | 2100 | | | | | | |
| | | - | , , | - | tgatgatcgg | - | 2160 | | | | | | |
| | | | | | atctcgaatc | | 2220 | | | | | | |
| | | | | | tgacaacttt | | 2280 | | | | | | |
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| | | | | | ccatcaaacc | | 60 | | | | | | |
| | | | | | gtctaagcac | | 120 | | | | | | |
| | | | | | ttgatctaaa | | 180 | | | | | | |
| | | | | | gggtagaaga | | 240 | | | | | | |
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| | | | | | tttgtgcact | gagcgatgtc | 360 | | | | | | |
| 150 | accatttcta | cctgccacgc | atcggcgaag | gttggaactt | ag | | 402 | | | | | | |
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| | <211> LENGT | | | | | | | | | | | | |
| | <212> TYPE: | | 1 | • | | | | | | | | | |
| | | NISM: Arabid | opsis thai | Lana | | | | | | | | | |
| | <400> SEQUE | | | | | | 60 | | | | | | |
| | | | | | ctcctttgat | | 60 120 | | | | | | |
| | | | | | tcttagacct | | 120 | | | | | | |
| | | | | | ccagtataga gttgtggcat | | 180 | | | | | | |
| | | | | | atgttcttgc | | 240 300 | | | | | | |
| 102 | aatttcaayg | citytyteea | aattotooya | ayaacttttg | acyclottyc | acttadycat | 300 | | | | | | |

RAW SEQUENCE LISTING DATE: 03/18/2002 PATENT APPLICATION: US/09/749,601A TIME: 14:45:17

Input Set : A:\00069.JHU.SEQ.TXT

Output Set: N:\CRF3\03182002\1749601A.raw

| 163 catacttota aattagagga tttcacagat ottttgaatt tgactaotta tggttttaga 164 ggagaagoot tgagototot otgtgoattg ggaaatotoa otgtggaaac aagaacaaag | 360 420 | | | | | | | | | | | | |
|--|------------|--|--|--|--|--|--|--|--|--|--|--|--|
| 165 aatgagccag ttgctacgct c | 441 | | | | | | | | | | | | |
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| 186 <213> ORGANISM: Arabidopsis thaliana | | | | | | | | | | | | | |
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| 189 Met Ile Asp Asp Ser Ser Leu Thr Ala Glu Met Glu Glu Glu Glu Ser | | | | | | | | | | | | | |
| 190 1 5 10 15 | | | | | | | | | | | | | |
| 191 Pro Ala Thr Thr Ile Val Pro Arg Glu Pro Pro Lys Ile Gln Arg Leu | | | | | | | | | | | | | |
| 192 20 25 30 | | | | | | | | | | | | | |
| 193 Glu Glu Ser Val Val Asn Arg Ile Ala Ala Gly Glu Val Ile Gln Arg | | | | | | | | | | | | | |
| 194 35 40 45 | | | | | | | | | | | | | |
| 195 Pro Val Ser Ala Val Lys Glu Leu Val Glu Asn Ser Leu Asp Ala Asp | | | | | | | | | | | | | |
| 196 50 55 60 | | | | | | | | | | | | | |
| 197 Ser Ser Ser Ile Ser Val Val Lys Asp Gly Gly Leu Lys Leu Ile | | | | | | | | | | | | | |
| 198 65 70 75 80 | | | | | | | | | | | | | |
| 199 Gln Val Ser Asp Asp Gly His Gly Ile Arg Arg Glu Asp Leu Pro Ile | | | | | | | | | | | | | |
| 200 85 90 95 | | | | | | | | | | | | | |
| 201 Leu Cys Glu Arg His Thr Thr Ser Lys Leu Thr Lys Phe Glu Asp Leu | | | | | | | | | | | | | |
| 202 100 105 110 | | | | | | | | | | | | | |
| 203 Phe Ser Leu Ser Ser Met Gly Phe Arg Gly Glu Ala Leu Ala Ser Met | | | | | | | | | | | | | |
| 204 115 120 125 | | | | | | | | | | | | | |
| 205 Thr Tyr Val Ala His Val Thr Val Thr Ile Thr Lys Gly Gln Ile | | | | | | | | | | | | | |
| 206 130 135 140 | | | | | | | | | | | | | |
| 207 His Gly Tyr Arg Val Ser Tyr Arg Asp Gly Val Met Glu His Glu Pro 208 145 150 155 160 | | | | | | | | | | | | | |
| | | | | | | | | | | | | | |
| 209 Lys Ala Cys Ala Ala Val Lys Gly Thr Gln Ile Met Val Glu Asn Leu 210 165 170 175 | | | | | | | | | | | | | |
| 211 Phe Tyr Asn Met Ile Ala Arg Arg Lys Thr Leu Gln Asn Ser Ala Asp | | | | | | | | | | | | | |
| 211 File Tyl Asii Met Tie Ald Ary Ary Lys Till Led Gill Asii Sel Ard Asp 212 180 185 190 | | | | | | | | | | | | | |
| 213 Asp Tyr Gly Lys Ile Val Asp Leu Leu Ser Arg Met Ala Ile His Tyr | | | | | | | | | | | | | |
| 214 195 200 205 | | | | | | | | | | | | | |
| 215 Asn Asn Val Ser Phe Ser Cys Arg Lys His Gly Ala Val Lys Ala Asp | | | | | | | | | | | | | |
| 216 210 215 220 | | | | | | | | | | | | | |
| 217 Val His Ser Val Val Ser Pro Ser Arg Leu Asp Ser Ile Arg Ser Val | | | | | | | | | | | | | |

RAW SEQUENCE LISTING DATE: 03/18/2002 PATENT APPLICATION: US/09/749,601A TIME: 14:45:17

Input Set : A:\00069.JHU.SEQ.TXT
Output Set: N:\CRF3\03182002\1749601A.raw

| | 225 | | | | | 230 | | | | | 235 | | | | | 240 |
|-----|----------|-------|------|------------|----------|------------|----------------|-------|------------|-----------|-----------|-------------|------------|------------------|----------|------------|
| 219 | Tyr | Gly | Val | Ser | Val | Ala | Lys | Asn | Leu | Met | Lys | Val | Glu | Val | Ser | Ser |
| 220 | | | | | 245 | | | | | 250 | | | | | 255 | |
| 221 | Cys | Asp | Ser | Ser | Gly | Cys | Thr | Phe | Asp | Met | Glu | Gly | Phe | Ile | Ser | Asn |
| 222 | - | - | | 260 | - | - | | | 265 | | | - | | 270 | | |
| | Ser | Asn | Tyr | Va 1 | Ala | Lvs | Lvs | Thr | | Leu | Val | Leu | Phe | | Asn | Asn |
| 224 | Der | non | 275 | vai | mu | цуз | цуз | 280 | 110 | пса | 141 | пси | 285 | 110 | ASII | пор |
| | 3 | T | | <i>α</i> 1 | 0 | C ~ ~ | 7 1 a | - | T | 7 | 7 7 ~ | T 1. | | T1_ | 37- 1 | m |
| | Arg | | Val | Gru | Cys | ser | | Leu | гуѕ | Arg | Ald | | GIU | TTE | Val | TAL |
| 226 | _ | 290 | | _ | _ | _ | 295 | _ | _ | _ | | 300 | _ | | _ | _ = |
| | | Ala | Thr | Leu | Pro | _ | Ala | Ser | Lys | Pro | | Val | Tyr | Met | Ser | Ile |
| 228 | 305 | | | | | 310 | | | | | 315 | | | | | 320 |
| 229 | Asn | Leu | Pro | Arg | | His | Val | Asp | Ile | Asn | Ile | His | Pro | \mathtt{Thr} | Lys | Lys |
| 230 | | | | | 325 | | | | | 330 | | | | | 335 | |
| 231 | Glu | Val | Ser | Leu | Leu | Asn | Gln | Glu | Ile | Ile | Ile | Glu | Met | Ile | Gln | Ser |
| 232 | | | | 340 | | | | | 345 | | | | | 350 | | |
| | Glu | Va l | Glu | Val | Lvs | Leu | Ara | Asn | Ala | Asn | Asp | Thr | Ara | Thr | Phe | Gln |
| 234 | | | 355 | | -1- | | 5 | 360 | | | | | 365 | | | 0 |
| | Clu | Cln | | Va I | Clu | Петъ | Tlo | | cor | Thr | LOU | mbr. | | Cln | Lys | Cor |
| | GIU | 370 | цуз | vai | Giu | тут | 375 | GIII | Ser | TIII | шeu | 380 | Ser | GIII | цуѕ | ser |
| 236 | | | D | **- 1 | a | a 1 | | D | a | 01 | 01 | | ml | 01 - | . | **- 3 |
| | | ser | Pro | vaı | ser | | ьуs | Pro | ser | GIY | | ьys | Thr | GIN | Lys | |
| | 385 | | | | | 390 | | | | | 395 | | | | | 400 |
| 239 | Pro | Val | Asn | Lys | Met | Val | Arg | Thr | Asp | | Ser | Asp | Pro | Ala | Gly | Arg |
| 240 | | | | | 405 | | | | | 410 | | | | | 415 | |
| 241 | Leu | His | Ala | Phe | Leu | Gln | Pro | Lys | Pro | Gln | Ser | Leu | Pro | Asp | Lys | Val |
| 242 | | | | 420 | | | | | 425 | | | | | 430 | | |
| 243 | Ser | Ser | Leu | Ser | Val | Val | Arg | Ser | Ser | Val | Arg | Gln | Arq | Arq | Asn | Pro |
| 244 | | | 435 | | | | | 440 | | | | | 445 | _ | | |
| 245 | Lvs | Glu | Thr | Ala | Asp | Leu | Ser | Ser | Val | Gln | Glu | Leu | Ile | Ala | Gly | Val |
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| | Asn | - | Cvs | Cvs | His | Pro | | Met | T.eu | Glu | Thr | | Δτα | Δgn | Cys | Thr |
| | 465 | DCI | CID | CID | 111.5 | 470 | O _T | 1100 | пса | OIu | 475 | Val | 1119 | 11511 | Cys | 480 |
| | | Wa 1 | C117 | Mot | ת 1 ת | | 7 an | Wa I | Dho | 717 | | Wa 1 | Cln | m.r.s | Asn | |
| | тут | var | СТУ | мес | | ASP | ASP | Val | PHE | | ьeu | val | GTII | тут | | 1111 |
| 250 | *** - | T | m | T | 485 | 3 | **- 1 | **- 1 | | 490 | a | T | a 1 | . | 495 | . |
| | HIS | Leu | Tyr | | Ата | Asn | vaı | vaı | | Leu | ser | гàг | Gru | | Met | Tyr |
| 252 | | _ | _ | 500 | | | _ | | 505 | | | _ | _ | 510 | | |
| | Gln | Gln | | Leu | Arg | Arg | Phe | | His | Phe | Asn | Ala | | Gln | Leu | Ser |
| 254 | | | 515 | | | | | 520 | | | | | 525 | | | |
| 255 | Asp | Pro | Ala | Pro | Leu | Ser | Glu | Leu | Ile | Leu | Leu | Ala | Leu | Lys | Glu | Glu |
| 256 | | 530 | | | | | 535 | | | | | 540 | | | | |
| 257 | Asp | Leu | Asp | Pro | Gly | Asn | Asp | Thr | Lys | Asp | Asp | Leu | Lys | Glu | Arg | Ile |
| 258 | | | _ | | _ | 550 | _ | | _ | _ | 555 | | _ | | _ | 560 |
| | | Glu | Met. | Asn | Thr | Glu | Leu | Leu | Lvs | Glu | Lvs | Ala | Glu | Met | Leu | |
| 260 | | | | | 565 | | | | ~1~ | 570 | -12 | | | | 575 | 0_0 |
| | Clu | Тиг | Dho | cor | | uic | T10 | λen | Sor | | λla | λαη | T OIL | Sor | Arg | LOU |
| | GIU | тут | FILE | | Vai | птэ | 116 | Аэр | | SET | Ala | ASII | Leu | | AIG | цец |
| 262 | D | 17- 7 | т1- | 580 | 3 | a 1 | m | m k | 585 Dec | 3 | We ± | 3 | 7 | 590 | D | a 1 |
| | Pro | val | | ren | ASP | GIN | туг | | PLO | ASP | met | Asp | | val | Pro | GLU |
| 264 | | _ | 595 | _ | _ | | _ | 600 | | _ • | _ | _ • | 605 | | _ | _ |
| | Phe | | Leu | Cys | Leu | Gly | | Asp | Val | Glu | Trp | | Asp | Glu _. | Lys | Ser |
| 266 | | 610 | | | | | 615 | | | | | 620 | | | | |

VERIFICATION SUMMARY

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